PTO/SB/08A (08-03)

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| Substitu               | te for form 1449A/P1     | ige,     | A            |                        | Complete if Known |
|                        |                          | . – .    | MOEMARY CEE  | Application Number     | 08/856,376        |
|                        | · · · · · <del>-</del> · |          | SCLOSURE     | Filing Date            | May 14, 1997      |
| STATEMENT BY APPLICANT |                          |          |              | First Named Inventor   | Chee, Mark        |
|                        |                          |          |              | Art Unit               | 1631              |
|                        | (use as many sh          | eets a   | s necessary) | Examiner Name          | Ardin Marschel    |
| Sheet                  | 1                        | of       | 3            | Attorney Docket Number | 018547-025010US   |

|                       | U.S. PATENT DOCUMENTS+   |  |                                |  |   |  |  |
|-----------------------|--------------------------|--|--------------------------------|--|---|--|--|
| Examiner<br>Initlals* | Cite<br>No. <sup>1</sup> | Document Number  Number Kind Code <sup>2</sup> ( <i>Il known</i> ) | Publication Date<br>MM-DD-YYYY | Name of Patentee or<br>Applicant of Cited Document | Pages, Columns, Lines, Where<br>Relevant Passages or Relevant<br>Figures Appear |  |  |

| FOREIGN PATENT DOCUMENTS |                          |                                |  |                                  |                                |   |  |    |
|--------------------------|--------------------------|--------------------------------|--|----------------------------------|--------------------------------|---|--|----|
| Examiner<br>Initials*    | Cite<br>No. <sup>1</sup> | Fore Country Code <sup>3</sup> | eign Patent Doo<br>Number <sup>4</sup> | Kind Code <sup>6</sup> (# known) | Publication Date<br>MM-DD-YYYY | Name of Patentee or<br>Applicant of Cited<br>Document | Pages, Columns, Lines,<br>Where Relevant<br>Passages or Relevant<br>Figures Appear | T⁵ |

Examiner Signature Andin Manshel Date Considered 9-17-04



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## INFORMATION DISCLOSURE STATEMENT BY APPLICANT

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| Complete if Known      |                 |  |  |  |
|------------------------|-----------------|--|--|--|
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| Examiner Name          | Ardin Marschel  |  |  |  |
| Attorney Docket Number | 018547-025010US |  |  |  |

|                     |              | NON PATENT LITERATURE DOCUMENTS   |     |
|---------------------|--------------|---|-----|
| Examiner Initials * | Cite<br>No.1 | Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published. | T 2 |
| AM                  | AA           | Ginther et al., "Identifying individuals by sequencing mitochondrial DNA from teeth," Nature Genetics, 2:135 (10/1992).   |     |
|                     | AB           | Greenberg et al., "Intraspecific nucleotide sequence variability surrounding the origin of replication in human mitochondrial DNA," Gene 21(1-2):33 (1983).   |     |
|                     | AC           | Howell et al., "Mitochondrial gene segregation in humans: is the bottleneck always narrow?" Human Genetics, 90:117 (1992).  |     |
|                     | AD           | Howell et al., "When does bilateral optic atrophy become Leber hereditary optic atrophy?" American Journal of Human Genetics, 53:959 (1993).  |     |
|                     | AE           | Hutchin et al., "A molecular basis for human hypersensitivity to aminoglycoside antibiotics," NAR 21(18):4174 (1993).   |     |
|                     | AF           | Ikebe et al., "Point mutations of mitochondrial genome in PArkinson's disease," Molecular Brain Research 28(2):281 (1995).  |     |
|                     | AG           | Isenberg and Moore, "Mitochondrial DNA Analysis at the FBI Laboratory," Forensic Science Communications, Vol. 1, No. 2 (7/1999).  |     |
|                     | АН           | Johns and Neufeld, "Pitfalls in the molecular genetic diagnosis of Leber hereditary optic neuropathy (LHON)," American Journal of Human Genetics, 53 (4):916 (1993).  |     |
|                     | Al           | Marzuki et al., "Normal variants of human mitochondrial DNA and translation products: building a reference data base," Human Genetics, 88 (2):139 (1991).   |     |
|                     | LA           | Mehta, et al., "A new genetic polymorphism in the 16S ribosomal RNA gene of human mitochondrial DNA," Annals of Human Genetics, 53 (Pt. 4):303 (1989).  |     |
|                     | AK           | Moraes, et al., "Two novel pathogenic mitochondrial DNA mutations affecting organelle number and protein synthesis. Is the tRNA Leu(UUR) gene an etiologic hot spot?" J. of Clinical Investigation, 92(6):2906 (1993).  |     |
|                     | AL           | Ozawa et al., "Distinct clustering of point mutations in mitochondrial DNA amoung patients with mitochondrial encephalomyopathies and with Parkinson's disease," BBRC, 176 (2):938 (1991).  |     |
|                     | АМ           | Ozawa et al., "Patients with idiopathic cardiomyopathy belong to the same mitochondrial gene family of Parkinson's disease and mitochondrial encephalomyopathy," BBRC 177(1):518 (1991).  |     |
|                     | AN           | Petruzzella et al., "Is a point mutation in the mitochondrial ND2 gene associated with Alzhelmer's disease?" BBRC 186:491 (1992).   |     |
|                     | AO           | Prezant et al., "Milochondrial ribosomal RNA mutation associated with both antibiotic-induced and non-syndromic deafness," Nature Genetics, 4 (3):289-294(1993)   |     |
| V                   | AP           | Reid et al., "Complete mtDNA sequence of a patient in a maternal pedigree with sensorineural deafness," Human Molecular Genetics, 3(8):1435 (1994).   |     |

| Examiner<br>Signature | Andre Marsly | Date<br>Considered | 9-17-04 |
|-----------------------|--------------|--------------------|---------|

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| AM                  | AQ           | Ruvolo et al., Mitochondrial COII sequences and modern huiman origins," Molecular Biology and Evolution, 10:1115 (1993).  |     |
|                     | AR           | Seneca et al., "Importance of sequence analysis in the NARP syndrome," J. Inherited Metabolic Disorders, 18 (1):97 (1995).  |     |
| 9                   | AS           | Tanaka and Ozawa, "Strand asymmetry in human mitochondrial DNA mutations," Genomics, 22(2):327 (1994).  |     |

| Examiner<br>Signature | Ardin  | Marsh | Date Considered | 9-17-04 |  |
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